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LOGINID:SSSPTA1626GMS

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

* * * * * Welcome to STN International * * * * *

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2 "Ask CAS" for self-help around the clock
NEWS 3 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/
USPAT2
NEWS 4 JAN 13 IPC 8 searching in IFIPAT, IFIUDB, and IFICDB
NEWS 5 JAN 13 New IPC 8 SEARCH, DISPLAY, and SELECT enhancements added to
INPADOC
NEWS 6 JAN 17 Pre-1988 INPI data added to MARPAT
NEWS 7 JAN 17 IPC 8 in the WPI family of databases including WPIFV
NEWS 8 JAN 30 Saved answer limit increased
NEWS 9 FEB 21 STN AnaVist, Version 1.1, lets you share your STN AnaVist
visualization results
NEWS 10 FEB 22 The IPC thesaurus added to additional patent databases on STN
NEWS 11 FEB 22 Updates in EPFULL; IPC 8 enhancements added
NEWS 12 FEB 27 New STN AnaVist pricing effective March 1, 2006
NEWS 13 FEB 28 MEDLINE/LMEDLINE reload improves functionality
NEWS 14 FEB 28 TOXCENTER reloaded with enhancements
NEWS 15 FEB 28 REGISTRY/ZREGISTRY enhanced with more experimental spectral
property data
NEWS 16 MAR 01 INSPEC reloaded and enhanced
NEWS 17 MAR 03 Updates in PATDPA; addition of IPC 8 data without attributes
NEWS 18 MAR 08 X.25 communication option no longer available after June 2006
NEWS 19 MAR 22 EMBASE is now updated on a daily basis
NEWS 20 APR 03 New IPC 8 fields and IPC thesaurus added to PATDPAFULL
NEWS 21 APR 03 Bibliographic data updates resume; new IPC 8 fields and IPC
thesaurus added in PCTFULL
NEWS 22 APR 04 STN AnaVist \$500 visualization usage credit offered
NEWS 23 APR 12 LINSPEC, learning database for INSPEC, reloaded and enhanced
NEWS 24 APR 12 Improved structure highlighting in FQHIT and QHIT display
in MARPAT
NEWS 25 APR 12 Derwent World Patents Index to be reloaded and enhanced during
second quarter; strategies may be affected

NEWS EXPRESS FEBRUARY 15 CURRENT VERSION FOR WINDOWS IS V8.01a,
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.
V8.0 AND V8.01 USERS CAN OBTAIN THE UPGRADE TO V8.01a AT
<http://download.cas.org/express/v8.0-Discover/>

NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS LOGIN Welcome Banner and News Items
NEWS IPC8 For general information regarding STN implementation of IPC 8

04/23/2006 10714066.trn

Enter NEWS followed by the item number or name to see news on that specific topic.

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* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 13:26:33 ON 23 APR 2006

=>

Uploading

THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE

Do you want to switch to the Registry File?

Choice (Y/n):

Switching to the Registry File...

Some commands only work in certain files. For example, the EXPAND command can only be used to look at the index in a file which has an index. Enter "HELP COMMANDS" at an arrow prompt (=>) for a list of commands which can be used in this file.

=> FILE REGISTRY

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'REGISTRY' ENTERED AT 13:26:49 ON 23 APR 2006

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STRUCTURE FILE UPDATES: 21 APR 2006 HIGHEST RN 881539-69-1

DICTIONARY FILE UPDATES: 21 APR 2006 HIGHEST RN 881539-69-1

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

*
* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added, *
* effective March 20, 2005. A new display format, IDERL, is now *
* available and contains the CA role and document type information. *
*

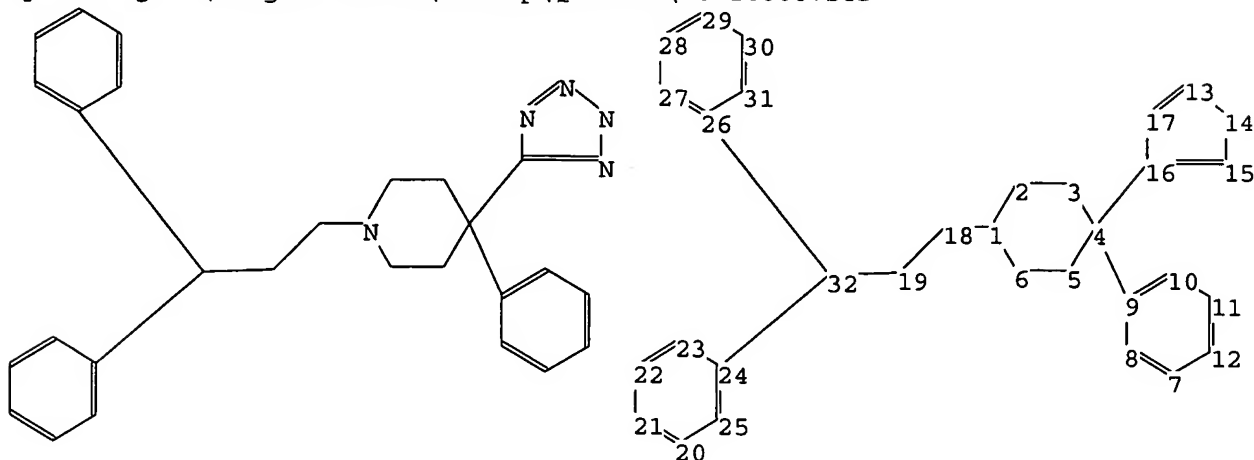
Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

=>

Uploading C:\Program Files\Stnexp\Queries\10714066.str



chain nodes :

18 19 32

ring nodes :

1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 20 21 22 23 24 25
26 27 28 29 30 31

chain bonds :

1-18 4-9 4-16 18-19 19-32 24-32 26-32

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 13-14 13-17
14-15 15-16 16-17 20-21 20-25 21-22 22-23 23-24 24-25 26-27 26-31 27-28
28-29 29-30 30-31

exact/norm bonds :

1-2 1-6 1-18 2-3 3-4 4-5 5-6 13-14 13-17 14-15 15-16 16-17

exact bonds :

4-9 4-16 18-19 19-32 24-32 26-32

normalized bonds :

7-8 7-12 8-9 9-10 10-11 11-12 20-21 20-25 21-22 22-23 23-24 24-25 26-27
26-31 27-28 28-29 29-30 30-31

isolated ring systems :

containing 1 : 7 : 13 : 20 : 26 :

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:CLASS 19:CLASS
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10714066.trn

Page 3

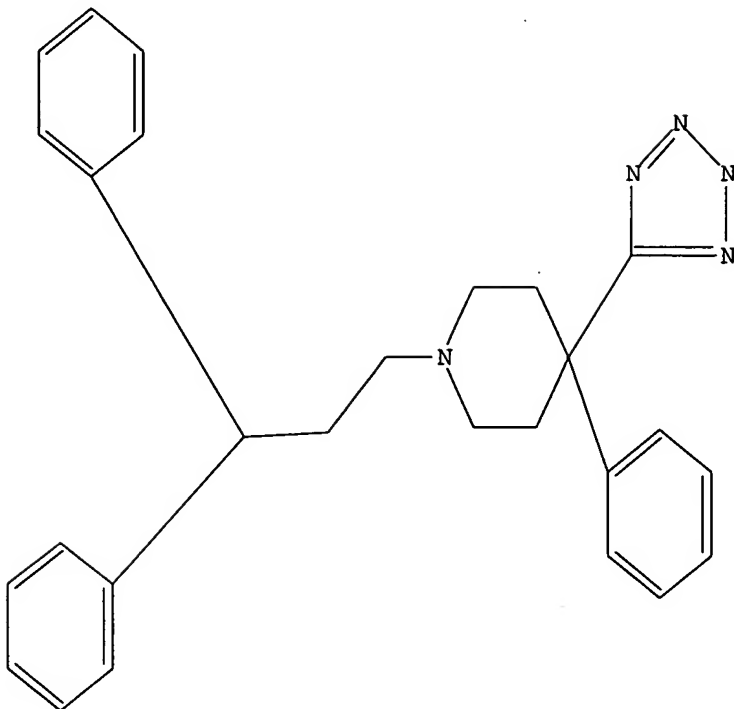
13:32

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 13:27:06 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 6 TO ITERATE

100.0% PROCESSED 6 ITERATIONS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 6 TO 266

PROJECTED ANSWERS: 2 TO 124

2 ANSWERS

L2 2 SEA SSS SAM L1

=> s l1 sss full

FULL SEARCH INITIATED 13:27:12 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 122 TO ITERATE

100.0% PROCESSED 122 ITERATIONS

11 ANSWERS

04/23/2006 10714066.trn

SEARCH TIME: 00.00.01

L3 ~~11 SEA SSS FUL L1~~

=> FIL HCAPLUS

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

166.94

167.15

FILE 'HCAPLUS' ENTERED AT 13:27:18 ON 23 APR 2006

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FILE COVERS 1907 - 23 Apr 2006 VOL 144 ISS 18

FILE LAST UPDATED: 21 Apr 2006 (20060421/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s l3

L4 ~~2 L3~~

=> d l4 ibib abs hitstr tot

L4 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:823687 HCAPLUS

DOCUMENT NUMBER: 143:211918

TITLE: Processes for preparation of 4-tetrazolyl-4-phenylpiperidines

INVENTOR(S): Brown, Kevin; Doyle, Timothy J.; Whitehead, John W. F.

PATENT ASSIGNEE(S): Euro-Celtique S.A., Luxembourg

SOURCE: PCT Int. Appl., 181 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005075455	A2	20050818	WO 2005-US3170	20050131
WO 2005075455	A3	20051215		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,

LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
 NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
 TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, SM
 RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
 AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
 EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
 RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
 MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.:

US 2004-540839P

P 20040130

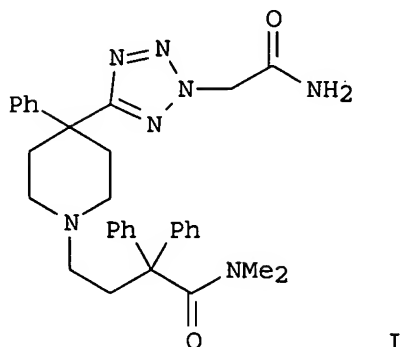
US 2004-552982P

P 20040311

OTHER SOURCE(S):

MARPAT 143:211918

GI



AB Processes are disclosed for the preparation of 4-tetrazolyl-4-phenylpiperidines. For instance, 4-cyano-4-phenylpiperidine•HCl is converted to 4-phenyl-4-(2H-tetrazol-5-yl)piperidine (water/dioxane, NaN₃, ZnBr₂, 90-100°, 24 h). This intermediate is reacted with (3,3-diphenyldihydrofuran-2-ylidene)dimethylammonium bromide (preparation given) (DMSO, DBU) to give N,N-dimethyl-2,2-diphenyl-4-[4-phenyl-4-(2H-tetrazol-5-yl)piperidin-1-yl]butanamide. This is treated with 2-chloroacetamide (DMSO, K₂CO₃, 60°, 4 h) to give I.

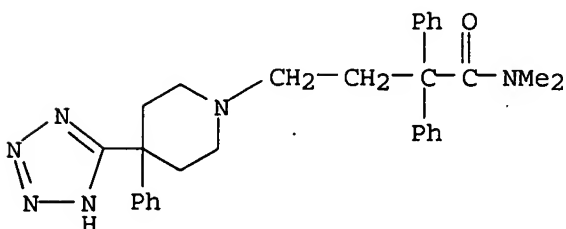
IT 697284-97-2P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(processes for preparation of 4-tetrazolyl-4-phenylpiperidines)

RN 697284-97-2 HCAPLUS

CN 1-Piperidinebutanamide, N,N-dimethyl- $\alpha,\alpha,4$ -triphenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)



IT 862287-78-3P 862287-79-4P 862287-93-2P

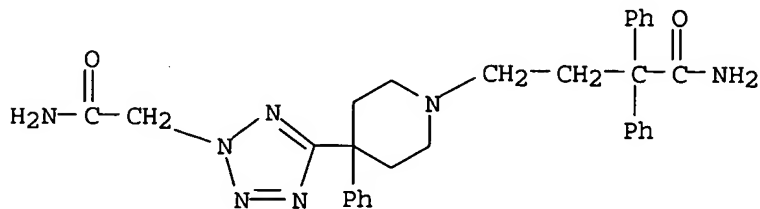
RL: SPN (Synthetic preparation); PREP (Preparation)

(processes for preparation of 4-tetrazolyl-4-phenylpiperidines)

04/23/2006 10714066.trn

RN 862287-78-3 HCAPLUS

CN 1-Piperidinebutanamide, 4-[2-(2-amino-2-oxoethyl)-2H-tetrazol-5-yl]-
 $\alpha,\alpha,4$ -triphenyl- (9CI) (CA INDEX NAME)



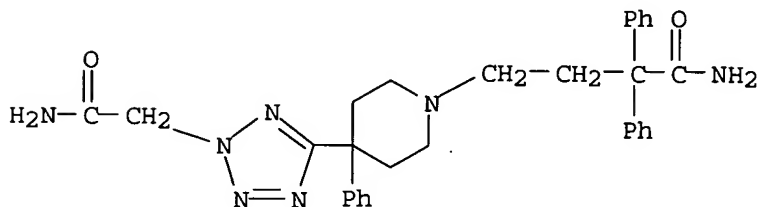
RN 862287-79-4 HCAPLUS

CN 1-Piperidinebutanamide, 4-[2-(2-amino-2-oxoethyl)-2H-tetrazol-5-yl]-
 $\alpha,\alpha,4$ -triphenyl-, monosulfamate (9CI) (CA INDEX NAME)

CM 1

CRN 862287-78-3

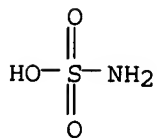
CMF C30 H33 N7 O2



CM 2

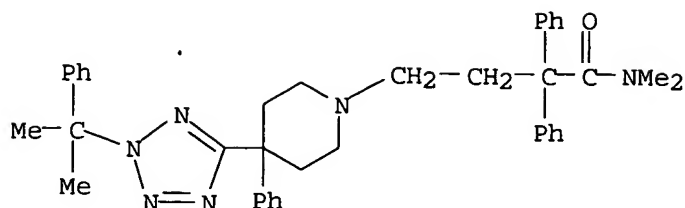
CRN 5329-14-6

CMF H3 N O3 S



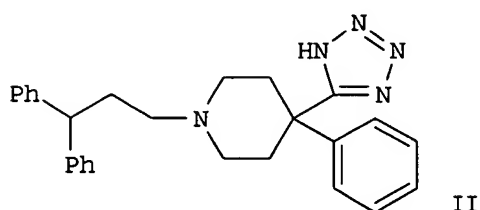
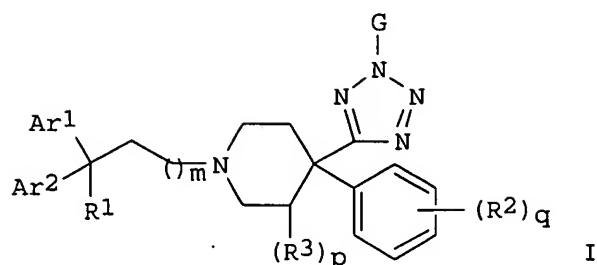
RN 862287-93-2 HCAPLUS

CN 1-Piperidinebutanamide, N,N-dimethyl-4-[2-(1-methyl-1-phenylethyl)-2H-
tetrazol-5-yl]- $\alpha,\alpha,4$ -triphenyl- (9CI) (CA INDEX NAME)



L4 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2004:453204 HCAPLUS
 DOCUMENT NUMBER: 141:7125
 TITLE: Preparation of 4-(tetrazol-5-yl)-4-phenylpiperidines
 for treating pain
 INVENTOR(S): Chen, Zhengming
 PATENT ASSIGNEE(S): Euro-Celtique, S.A., Luxembourg
 SOURCE: PCT Int. Appl., 136 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004046132	A1	20040603	WO 2003-US36742	20031117
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2004152689	A1	20040805	US 2003-714066	20031113
CA 2506242	AA	20040603	CA 2003-2506242	20031117
AU 2003294313	A1	20040615	AU 2003-294313	20031117
EP 1562932	A1	20050817	EP 2003-789796	20031117
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
BR 2003016305	A	20050927	BR 2003-16305	20031117
NO 2005002894	A	20050614	NO 2005-2894	20050614
PRIORITY APPLN. INFO.:			US 2002-427381P	P 20021118
			US 2003-460278P	P 20030403
			US 2003-488488P	P 20030717
			US 2003-714066	A 20031113
			WO 2003-US36742	W 20031117
OTHER SOURCE(S):		MARPAT 141:7125		
GI				



AB Title compds. I [Ar1 = cycloalkyl, Ph, naphthyl, etc.; Ar2 = Ph, naphthyl, anthryl, etc.; G = H, L(CH2)n-carboxy, etc.; L = CO, SO2, SO; R1 = H, carboxamido, etc.; R2-3 = halo, alkyl, alkoxy, etc.; m = 0-4; n = 1-4; p = 0-1; q = 0-3] are prepared For instance, 3,3-diphenyl-1-bromopropane is reacted with 4-cyano-4-phenylpiperidine (DMF, DIEA, 80°) and the alkylation product is converted to II (PhMe, TMSN3, Bu2SnO, reflux). Example compds. bind to μ -opioid and ORL1-receptors and are useful for the treatment of pain.

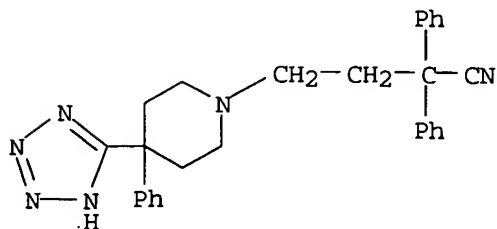
IT **697284-93-8P 697284-97-2P**

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of 4-(tetrazol-5-yl)-4-phenylpiperidines for treating pain)

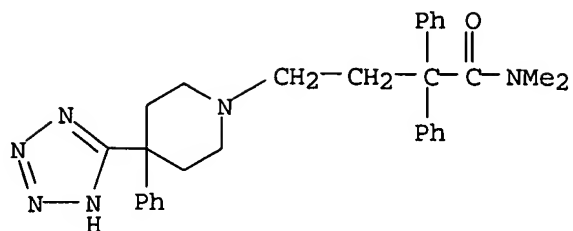
RN 697284-93-8 HCAPLUS

CN 1-Piperidinebutanenitrile, $\alpha,\alpha,4$ -triphenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)



RN 697284-97-2 HCAPLUS

CN 1-Piperidinebutanamide, N,N-dimethyl- $\alpha,\alpha,4$ -triphenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)



IT 697284-92-7P 697284-94-9P 697284-95-0P

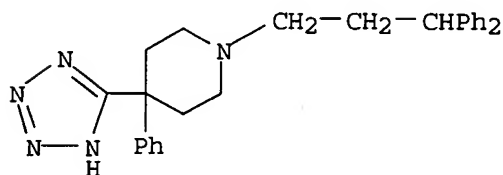
697284-96-1P 697284-98-3P 697284-99-4P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of 4-(tetrazol-5-yl)-4-phenylpiperidines for treating pain)

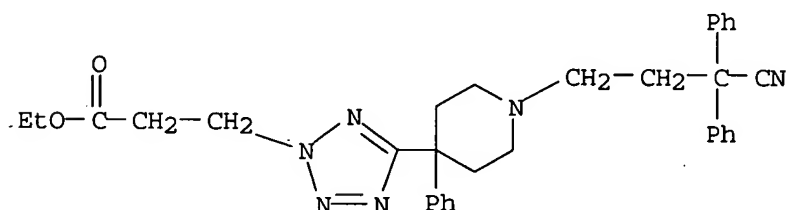
RN 697284-92-7 HCAPLUS

CN Piperidine, 1-(3,3-diphenylpropyl)-4-phenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)



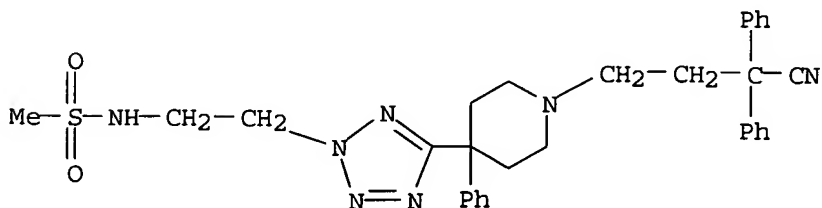
RN 697284-94-9 HCAPLUS

CN 2H-Tetrazole-2-propanoic acid, 5-[1-(3-cyano-3,3-diphenylpropyl)-4-phenyl-4-piperidinyl]-, ethyl ester (9CI) (CA INDEX NAME)



RN 697284-95-0 HCAPLUS

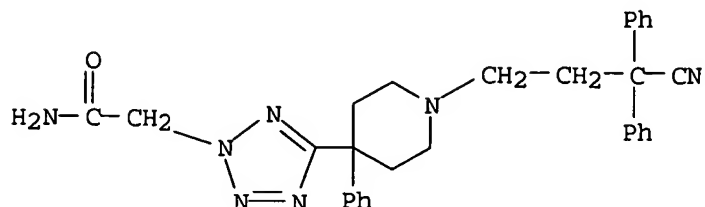
CN Methanesulfonamide, N-[2-[5-[1-(3-cyano-3,3-diphenylpropyl)-4-phenyl-4-piperidinyl]-2H-tetrazol-2-yl]ethyl]- (9CI) (CA INDEX NAME)



RN 697284-96-1 HCAPLUS

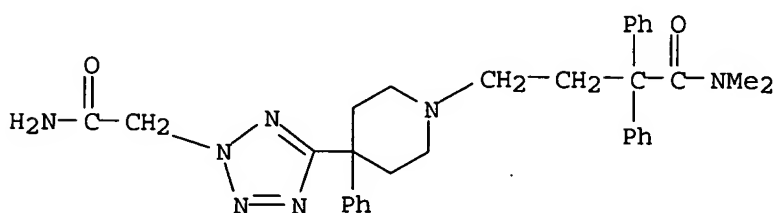
CN 2H-Tetrazole-2-acetamide, 5-[1-(3-cyano-3,3-diphenylpropyl)-4-phenyl-4-

piperidinyl]- (9CI) (CA INDEX NAME)



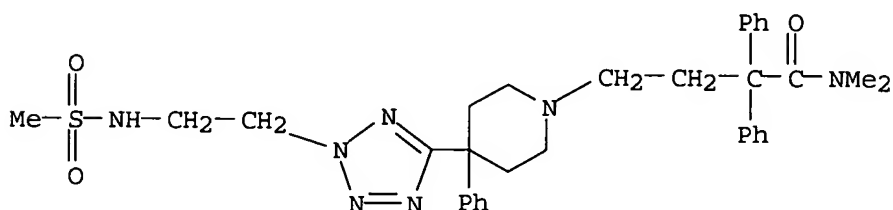
RN 697284-98-3 HCAPLUS

CN 1-Piperidinebutanamide, 4-[2-(2-amino-2-oxoethyl)-2H-tetrazol-5-yl]-N,N-dimethyl-α,α,4-triphenyl- (9CI) (CA INDEX NAME)



RN 697284-99-4 HCAPLUS

CN 1-Piperidinebutanamide, N,N-dimethyl-4-[2-[2-[(methylsulfonyl)amino]ethyl]-2H-tetrazol-5-yl]-α,α,4-triphenyl- (9CI) (CA INDEX NAME)



=> FIL REGISTRY

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
20.34	187.49

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
ENTRY	SESSION
-1.50	-1.50

CA SUBSCRIBER PRICE

FILE 'REGISTRY' ENTERED AT 13:29:27 ON 23 APR 2006

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04/23/2006 10714066.trn

STRUCTURE FILE UPDATES: 21 APR 2006 HIGHEST RN 881539-69-1
DICTIONARY FILE UPDATES: 21 APR 2006 HIGHEST RN 881539-69-1

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

Please note that search-term pricing does apply when
conducting SmartSELECT searches.

*
* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added, *
* effective March 20, 2005. A new display format, IDERL, is now *
* available and contains the CA role and document type information. *
*

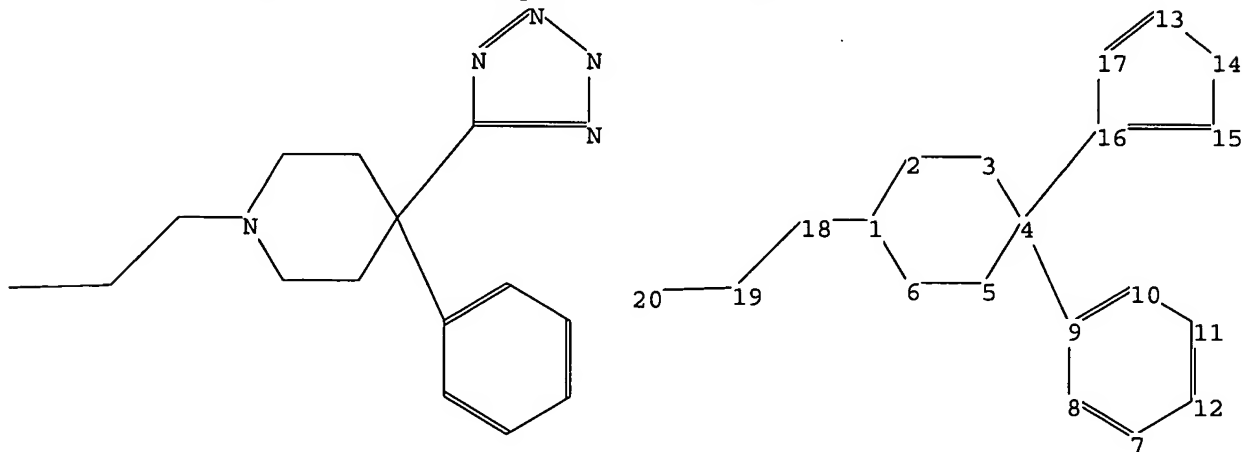
Structure search iteration limits have been increased. See HELP SLIMITS
for details.

REGISTRY includes numerically searchable data for experimental and
predicted properties as well as tags indicating availability of
experimental property data in the original document. For information
on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

=>

Uploading C:\Program Files\Stnexp\Queries\10714066a.str



chain nodes :

18 19 20

ring nodes :

1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17

chain bonds :

1-18 4-9 4-16 18-19 19-20

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 13-14 13-17
14-15 15-16 16-17

exact/norm bonds :

1-2 1-6 1-18 2-3 3-4 4-5 5-6 13-14 13-17 14-15 15-16 16-17

04/23/2006 10714066.trn

exact bonds :

4-9 4-16 18-19 19-20

normalized bonds :

7-8 7-12 8-9 9-10 10-11 11-12

isolated ring systems :

containing 1 : 7 : 13 :

Match level :

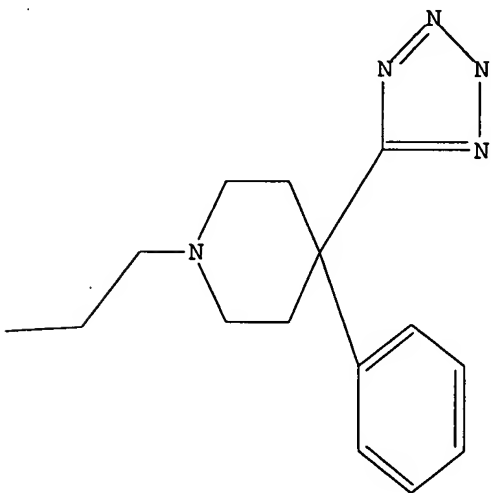
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:CLASS 19:CLASS
20:CLASS

L5 STRUCTURE UPLOADED

=> d 15

L5 HAS NO ANSWERS

L5 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 15

SAMPLE SEARCH INITIATED 13:29:51 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 6 TO ITERATE

100.0% PROCESSED 6 ITERATIONS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS: 6 TO 266

PROJECTED ANSWERS: 3 TO 163

3 ANSWERS

L6 3 SEA SSS SAM L5

04/23/2006 10714066.trn

=> s 15 sss full
FULL SEARCH INITIATED 13:29:59 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 122 TO ITERATE

100.0% PROCESSED 122 ITERATIONS 17 ANSWERS
SEARCH TIME: 00.00.01

L7 17 SEA SSS FUL L5

=> FIL HCAPLUS		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	166.94	354.43
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	0.00	-1.50

FILE 'HCAPLUS' ENTERED AT 13:30:04 ON 23 APR 2006
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FILE COVERS 1907 - 23 Apr 2006 VOL 144 ISS 18
FILE LAST UPDATED: 21 Apr 2006 (20060421/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 117
L17 NOT FOUND
The L-number entered has not been defined in this session, or it has been deleted. To see the L-numbers currently defined in this session, enter DISPLAY HISTORY at an arrow prompt (=>).

=> s 17
L8 4 L7

=> d 18 ibib abs hitstr tot

L8 ANSWER 1 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2005:823687 HCAPLUS
DOCUMENT NUMBER: 143:211918
TITLE: Processes for preparation of 4-tetrazolyl-4-phenylpiperidines
INVENTOR(S): Brown, Kevin; Doyle, Timothy J.; Whitehead, John W. F.

PATENT ASSIGNEE(S): Euro-Celtique S.A., Luxembourg
 SOURCE: PCT Int. Appl., 181 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

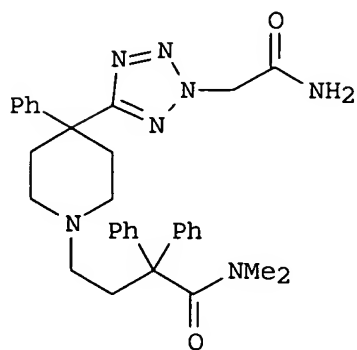
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005075455	A2	20050818	WO 2005-US3170	20050131
WO 2005075455	A3	20051215		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, SM

RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2004-540839P P 20040130
 US 2004-552982P P 20040311

OTHER SOURCE(S): MARPAT 143:211918
 GI



I

AB Processes are disclosed for the preparation of 4-tetrazolyl-4-phenylpiperidines. For instance, 4-cyano-4-phenylpiperidine•HCl is converted to 4-phenyl-4-(2H-tetrazol-5-yl)piperidine (water/dioxane, NaN₃, ZnBr₂, 90-100°, 24 h). This intermediate is reacted with (3,3-diphenyldihydrofuran-2-ylidene)dimethylammonium bromide (preparation given) (DMSO, DBU) to give N,N-dimethyl-2,2-diphenyl-4-[4-phenyl-4-(2H-tetrazol-5-yl)piperidin-1-yl]butyramide. This is treated with 2-chloroacetamide (DMSO, K₂CO₃, 60°, 4 h) to give I.

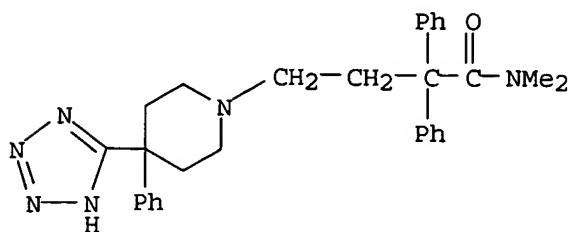
IT 697284-97-2P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(processes for preparation of 4-tetrazolyl-4-phenylpiperidines)

RN 697284-97-2 HCAPLUS

CN 1-Piperidinebutanamide, N,N-dimethyl- $\alpha,\alpha,4$ -triphenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)

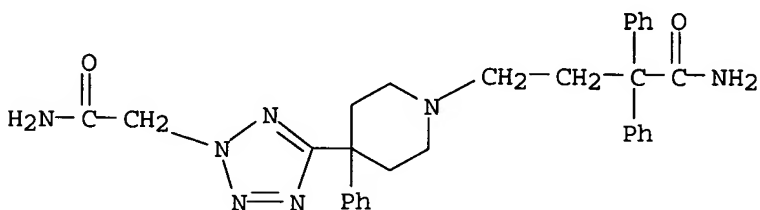


IT 862287-78-3P 862287-79-4P 862287-93-2P

RL: SPN (Synthetic preparation); PREP (Preparation)
(processes for preparation of 4-tetrazolyl-4-phenylpiperidines)

RN 862287-78-3 HCAPLUS

CN 1-Piperidinebutanamide, 4-[2-(2-amino-2-oxoethyl)-2H-tetrazol-5-yl]- $\alpha,\alpha,4$ -triphenyl- (9CI) (CA INDEX NAME)



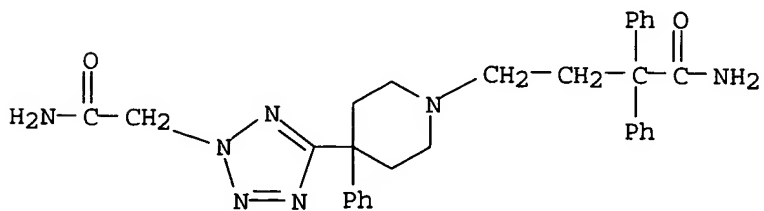
RN 862287-79-4 HCAPLUS

CN 1-Piperidinebutanamide, 4-[2-(2-amino-2-oxoethyl)-2H-tetrazol-5-yl]- $\alpha,\alpha,4$ -triphenyl-, monosulfamate (9CI) (CA INDEX NAME)

CM 1

CRN 862287-78-3

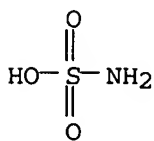
CMF C30 H33 N7 O2



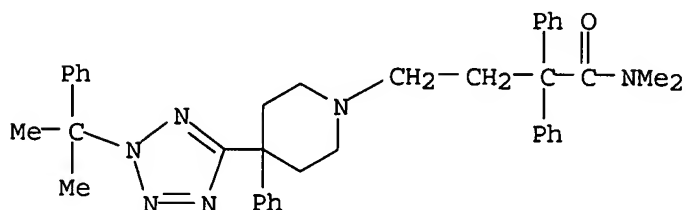
CM 2

CRN 5329-14-6

CMF H3 N O3 S



RN 862287-93-2 HCAPLUS

CN 1-Piperidinebutanamide, N,N-dimethyl-4-[2-(1-methyl-1-phenylethyl)-2H-tetrazol-5-yl]- $\alpha,\alpha,4$ -triphenyl- (9CI) (CA INDEX NAME)

L8 ANSWER 2 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:453204 HCAPLUS

DOCUMENT NUMBER: 141:7125

TITLE: Preparation of 4-(tetrazol-5-yl)-4-phenylpiperidines for treating pain

INVENTOR(S): Chen, Zhengming

PATENT ASSIGNEE(S): Euro-Genetique, S.A., Luxembourg

SOURCE: PCT Int. Appl., 136 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

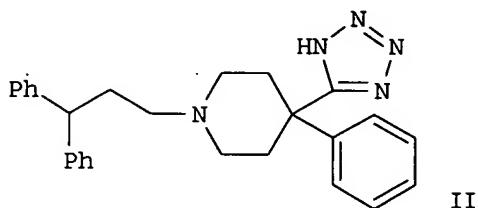
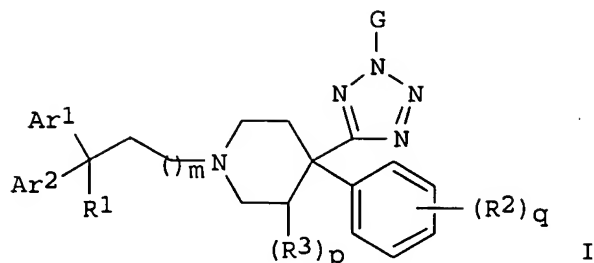
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004046132	A1	20040603	WO 2003-US36742	20031117
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2004152689	A1	20040805	US 2003-714066	20031113
CA 2506242	AA	20040603	CA 2003-2506242	20031117
AU 2003294313	A1	20040615	AU 2003-294313	20031117
EP 1562932	A1	20050817	EP 2003-789796	20031117
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
BR 2003016305	A	20050927	BR 2003-16305	20031117
NO 2005002894	A	20050614	NO 2005-2894	20050614
PRIORITY APPLN. INFO.:			US 2002-427381P	P 20021118

US 2003-460278P	P 20030403
US 2003-488488P	P 20030717
US 2003-714066	A 20031113
WO 2003-US36742	W 20031117

OTHER SOURCE(S): MARPAT 141:7125
GI



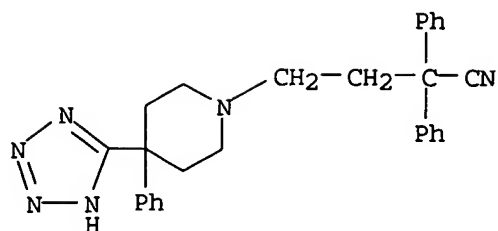
AB Title compds. I [Ar1 = cycloalkyl, Ph, naphthyl, etc.; Ar2 = Ph, naphthyl, anthryl, etc.; G = H, L(CH2)n-carboxy, etc.; L = CO, SO2, SO; R1 = H, carboxamido, etc.; R2-3 = halo, alkyl, alkoxy, etc.; m = 0-4; n = 1-4; p = 0-1; q = 0-3] are prepared For instance, 3,3-diphenyl-1-bromopropane is reacted with 4-cyano-4-phenylpiperidine (DMF, DIEA, 80°) and the alkylation product is converted to II (PhMe, TMSN3, Bu2SnO, reflux). Example compds. bind to μ -opioid and ORL1-receptors and are useful for the treatment of pain.

IT 697284-93-8P 697284-97-2P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
(preparation of 4-(tetrazol-5-yl)-4-phenylpiperidines for treating pain)

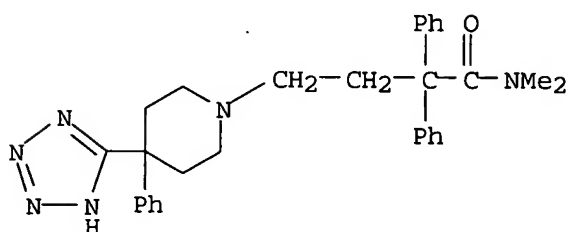
RN 697284-93-8 HCAPLUS

CN 1-Piperidinebutanenitrile, $\alpha,\alpha,4$ -triphenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)



RN 697284-97-2 HCAPLUS

CN 1-Piperidinebutanamide, N,N-dimethyl- $\alpha,\alpha,4$ -triphenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)



IT 697284-92-7P 697284-94-9P 697284-95-0P

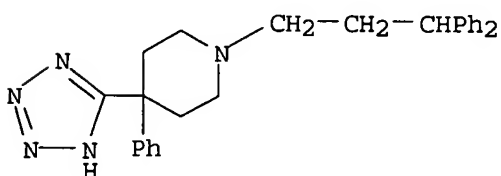
697284-96-1P 697284-98-3P 697284-99-4P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of 4-(tetrazol-5-yl)-4-phenylpiperidines for treating pain)

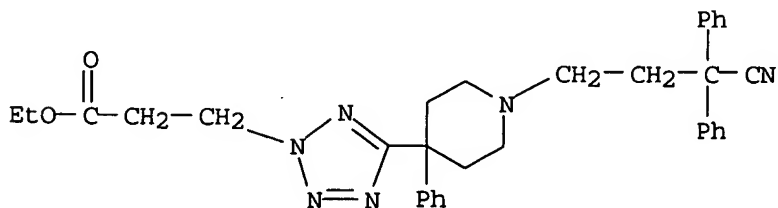
RN 697284-92-7 HCAPLUS

CN Piperidine, 1-(3,3-diphenylpropyl)-4-phenyl-4-(1H-tetrazol-5-yl)- (9CI) (CA INDEX NAME)



RN 697284-94-9 HCAPLUS

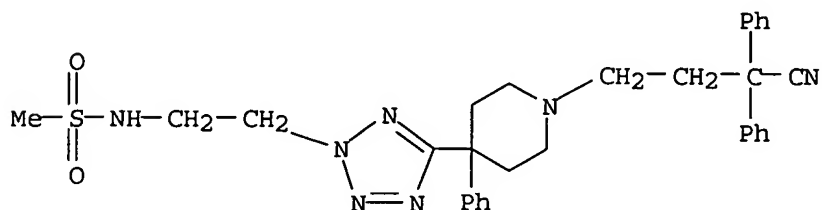
CN 2H-Tetrazole-2-propanoic acid, 5-[1-(3-cyano-3,3-diphenylpropyl)-4-phenyl-4-piperidiny]-, ethyl ester (9CI) (CA INDEX NAME)



04/23/2006 10714066.trn

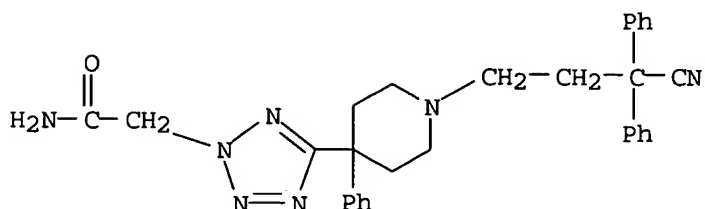
RN 697284-95-0 HCAPLUS

CN Methanesulfonamide, N-[2-[5-[1-(3-cyano-3,3-diphenylpropyl)-4-phenyl-4-piperidinyl]-2H-tetrazol-2-yl]ethyl]- (9CI) (CA INDEX NAME)



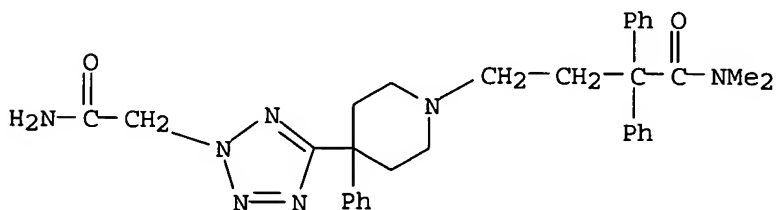
RN 697284-96-1 HCAPLUS

CN 2H-Tetrazole-2-acetamide, 5-[1-(3-cyano-3,3-diphenylpropyl)-4-phenyl-4-piperidinyl]- (9CI) (CA INDEX NAME)



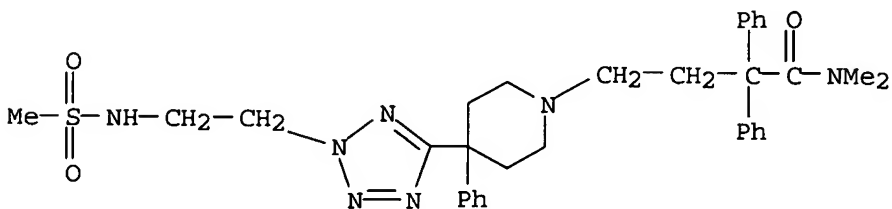
RN 697284-98-3 HCAPLUS

CN 1-Piperidinebutanamide, 4-[2-(2-amino-2-oxoethyl)-2H-tetrazol-5-yl]-N,N-dimethyl- $\alpha,\alpha,4$ -triphenyl- (9CI) (CA INDEX NAME)



RN 697284-99-4 HCAPLUS

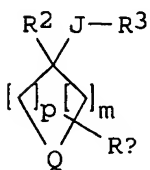
CN 1-Piperidinebutanamide, N,N-dimethyl-4-[2-[2-[(methylsulfonyl)amino]ethyl]-2H-tetrazol-5-yl]- $\alpha,\alpha,4$ -triphenyl- (9CI) (CA INDEX NAME)



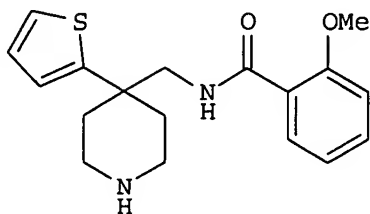
L8 ANSWER 3 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2003:855758 HCAPLUS

DOCUMENT NUMBER: 139:364829
 TITLE: Preparation of heterocyclo inhibitors of potassium channel function
 INVENTOR(S): Lloyd, John; Jeon, Yoon T.; Finlay, Heather; Yan, Lin; Beaudoin, Serge; Gross, Michael F.
 PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA; Icagen, Inc.
 SOURCE: PCT Int. Appl., 330 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003088908	A2	20031030	WO 2003-US11807	200310416
WO 2003088908	A3	20040527		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
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EP 1501467	A2	20050202	EP 2003-719792	20030416
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK			
JP 2005529114	T2	20050929	JP 2003-585661	20030416
NO 2004004351	A	20041013	NO 2004-4351	20041013
PRIORITY APPLN. INFO.:			US 2002-374279P	P 20020419
			WO 2003-US11807	W 20030416
OTHER SOURCE(S):	MARPAT 139:364829			
GI				



I



II

AB The title compds. [I; m, p = 0-3 (provided that the sum of m and p is at least 2); Q = NR1, O, S, SO, SO2; R1 = H, C(:W)NR6R7, SO2NR6R7, OCONR6R7, etc.; R2 = heteroaryl, heteroarylalkyl, aryl, etc.; J = a bond, alkylene; R3 = R5, OR5, SO2R5, etc.; R5 = CN, heteroaryl, aryl, etc.; R6, R7 = H, alkyl, OH, etc.; W = (un)substituted NH, N(CO2H), N(CN), N(SO2H), CH(NO2); Rx = H, alkyl, hydroxyalkyl, aryl, etc.], useful as inhibitors of potassium channel function (especially inhibitors of the Kv1 subfamily of voltage gated K⁺ channels, especially inhibitors Kv1.5 which has been linked to the ultra-rapidly activating delayed rectifier K⁺ current IK_{ur}) in the prevention and treatment of arrhythmia and IK_{ur}-associated conditions, were

prepared E.g., a multi-step synthesis of II [starting from bis(2-chloroethyl)amine], was given. Pharmaceutical composition comprising the compound I is claimed.

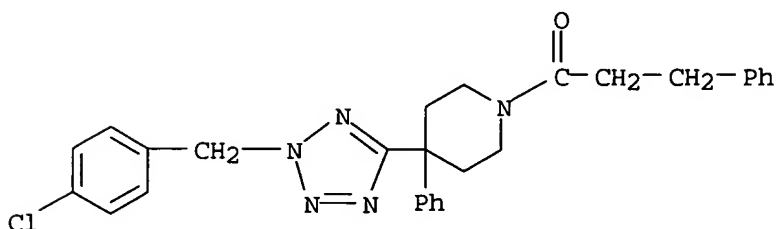
IT 619291-49-5P 619291-50-8P 619291-51-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of substituted piperidines as inhibitors of potassium channel function)

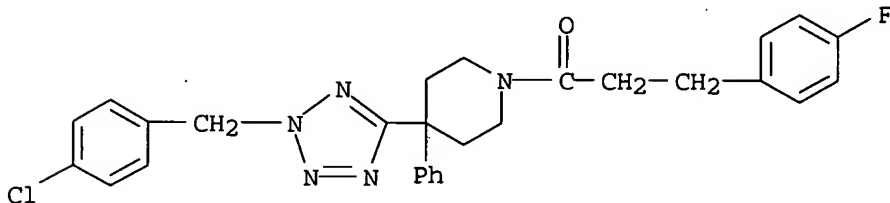
RN 619291-49-5 HCAPLUS

CN Piperidine, 4-[2-[(4-chlorophenyl)methyl]-2H-tetrazol-5-yl]-1-(1-oxo-3-phenylpropyl)-4-phenyl- (9CI) (CA INDEX NAME)



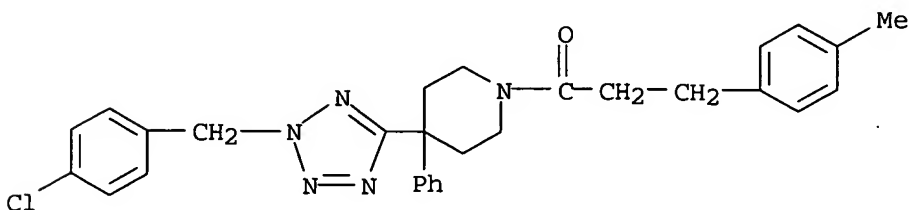
RN 619291-50-8 HCAPLUS

CN Piperidine, 4-[2-[(4-chlorophenyl)methyl]-2H-tetrazol-5-yl]-1-[3-(4-fluorophenyl)-1-oxopropyl]-4-phenyl- (9CI) (CA INDEX NAME)



RN 619291-51-9 HCAPLUS

CN Piperidine, 4-[2-[(4-chlorophenyl)methyl]-2H-tetrazol-5-yl]-1-[3-(4-methylphenyl)-1-oxopropyl]-4-phenyl- (9CI) (CA INDEX NAME)



L8 ANSWER 4 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN

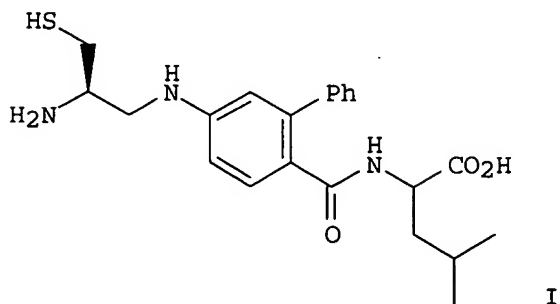
ACCESSION NUMBER: 2000:68365 HCAPLUS

DOCUMENT NUMBER: 132:122932

TITLE: Preparation of peptides, peptidomimetics, and nonpeptides as medical and agrochemical antifungals.

INVENTOR(S): Bergnes, Gustave; Berlin, Vivian; Come, Jon; Kluge, Arthur; Murthi, Krishna; Pal, Kollol
 PATENT ASSIGNEE(S): Mitotix, Inc., USA
 SOURCE: PCT Int. Appl., 287 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 200003743	A2	20000127	WO 1999-US16146	19990715
WO 200003743	A3	20010201		
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RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6423519	B1	20020723	US 1998-172845	19981015
CA 2335381	AA	20000127	CA 1999-2335381	19990715
AU 9951075	A1	20000207	AU 1999-51075	19990715
EP 1096925	A2	20010509	EP 1999-935639	19990715
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2002520372	T2	20020709	JP 2000-559877	19990715
PRIORITY APPLN. INFO.:			US 1998-115846	A 19980715
			US 1998-172845	A 19981015
			WO 1999-US16146	W 19990715
OTHER SOURCE(S):		MARPAT 132:122932		
GI				



AB A method for inhibiting the growth of a fungal pathogen comprises contacting the pathogen with a compound, e.g., (R70)2NCH[(CH2)nR]C(Xa)NHCHR72C(Xb)NHCHR73C(Xc)NHCHR10CO2R11 [Xa, Xb, Xc = O, H2; R = SR1, SOR111, SO2R111; R1 = H, alkyl, alkenyl, aryl, acyl; R10 = alkyl, alkenyl, alkynyl, aryl, cycloalkyl, hydroxyalkyl, amino acid sidechain, etc.; R11 = H, blocking group, pharmaceutically acceptable salt; R10R11 = atoms to

form 5-7 membered ring; R111 = alkyl, alkenyl, (CH₂)_mR₇; R₇₀ = H, alkyl, alkenyl, alkynyl, aryl, acyl, amino acid sidechain, etc.; R₇₂, R₇₃ = H, alkyl, aryl, heteroaryl, amino acid sidechain, (CH₂)_mR₇, etc.; m, n = 0-4], which inhibits prenyl transferase activity with MIC₅₀<25 µg/mL. Thus, title compound (I) (solution phase preparation given) inhibited GGTase

with

IC₅₀<10 nM.

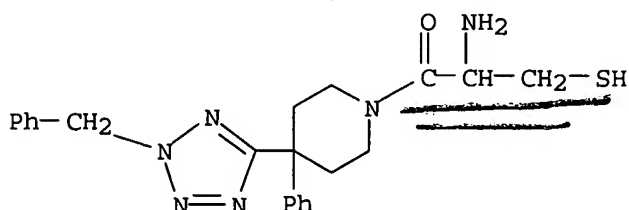
IT 256369-69-4 256369-76-3 256369-84-3

RL: AGR (Agricultural use); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(preparation of peptides, peptidomimetics, and nonpeptides as medical and agrochem. antifungals)

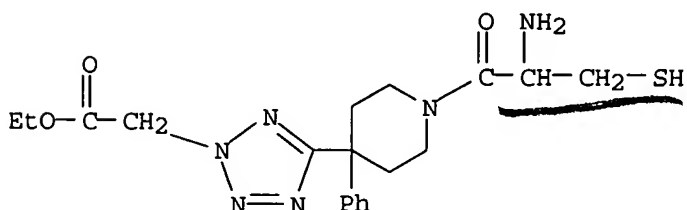
RN 256369-69-4 HCAPLUS

CN Piperidine, 1-(2-amino-3-mercapto-1-oxopropyl)-4-phenyl-4-[2-(phenylmethyl)-2H-tetrazol-5-yl]- (9CI) (CA INDEX NAME)



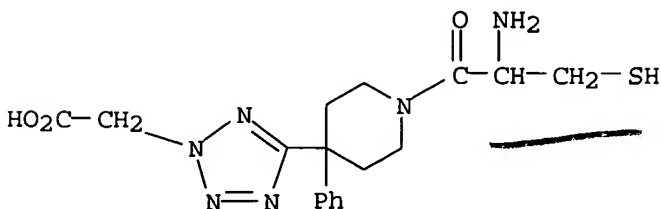
RN 256369-76-3 HCAPLUS

CN 2H-Tetrazole-2-acetic acid, 5-[1-(2-amino-3-mercapto-1-oxopropyl)-4-phenyl-4-piperidinyl]-, ethyl ester (9CI) (CA INDEX NAME)



RN 256369-84-3 HCAPLUS

CN 2H-Tetrazole-2-acetic acid, 5-[1-(2-amino-3-mercapto-1-oxopropyl)-4-phenyl-4-piperidinyl]- (9CI) (CA INDEX NAME)



=> log y

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

04/23/2006 10714066.trn

FULL ESTIMATED COST	ENTRY 25.50	SESSION 379.93
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	-3.00	-4.50

STN INTERNATIONAL LOGOFF AT 13:31:11 ON 23 APR 2006